CLAIMS

1. A method for treating chronic pain, said method comprising administering to a subject in need of such treatment a composition comprising a MEK inhibitor selected from: a compound of formula (I):

$$R_1 \xrightarrow{W} X \xrightarrow{R_4} R_3$$

$$R_2 \qquad (I)$$

10

5

W is one of the following formulae (i) - (xiii):

 X_2 X_1 X_2 X_1

O X₁ N (xv)

O X₁ (xvi)

N-N N N (xvii)

15

X₁ is O, S, or NR_F;

X₂ is OH, SH, or NHR_E;

each of R_E and R_F is H or C ₁₋₄ alkyl;

5

each of R_1 and R_2 is independently selected from H, F, NO₂, Br and Cl; R_1 can also be $SO_2NR_GR_H$, or R_1 and R_2 together with the benzene ring to which they are attached constitute an indole, isoindole, benzofuran, benzothiophene, indazole, benzimidazole, or benzthioazole;

10

R₃ is H or F;

each of R_G, R_H, and R₄ is independently selected from H, Cl and CH₃;

15

R₅ is H or C ₃₋₄ alkyl; and

wherein each hydrocarbon radical above is optionally substituted with between 1 and 3 substituents independently selected from halo, hydroxyl, amino, (amino)sulfonyl, and NO₂; and

20

wherein each heterocyclic radical above is optionally substituted with between 1 and 3 substituents independently selected from halo, C $_{3-4}$ alkyl, C $_{3-6}$ cycloalkyl, C $_{3-4}$ alkenyl, C $_{3-4}$ alkynyl, phenyl, hydroxyl, amino, (amino)sulfonyl, and NO $_2$, wherein each substituent alkyl, cycloalkyl, alkenyl, alkynyl or phenyl is in turn optionally substituted with between 1 and 2 substituents independently selected from halo, C $_{1-2}$ alkyl, hydroxyl, amino, and NO $_2$;

or a pharmaceutically acceptable salt or C $_{\mbox{\scriptsize 1-8}}$ ester thereof.

30

25

10

25

10031149 "060305 PCT/US00/18346

- 2. The method of claim 1, wherein said chronic pain is selected from neuropathic pain, idiopathic pain, and pain associated with chronic alcoholism, vitamin deficiency, uremia, or hypothyroidism.
- 3. The method of claim 2, wherein said chronic pain is a type of neuropathic pain.
 - 4. The method of claim 3, wherein said neuropathic pain is associated with one of the following: inflammation, postoperative pain, phantom limb pain, burn pain, gout, trigeminal neuralgia, acute herpetic and postherpetic pain, causalgia, diabetic neuropathy, plexus avulsion, neuroma, vasculitis, viral infection, crush injury, constriction injury, tissue injury, limb amputation, post-operative pain, arthritis pain, and any other nerve injury between the peripheral nervous system and the central nervous system, inclusively.
- 5. The method of claim 2, wherein said chronic pain is associated with chronic alcoholism, vitamin deficiency, uremia, or hypothyroidism.
 - 6. The method of claim 2, wherein said chronic pain is associated with idiopathic pain.
- 7. The method of claim 1, wherein said chronic pain is associated with inflammation.
 - 8. The method of claim 1, wherein said chronic pain is associated with arthritis.
 - 9. The method of claim 1, wherein said chronic pain is associated with post-operative pain.
- 30 10. The method of claim 1, wherein R_1 is bromo or chloro.

- 11. A method of claim 1, wherein R_2 is fluoro.
- 12. A method of claim 1, wherein R₃ is H.

5

- 13. A method of claim 12, wherein each of R_2 and R_3 is H.
- 14. A method of claim 1, wherein each of R_2 and R_3 is fluoro.
- 10 15. A method of claim 14, wherein R_1 is bromo.
 - 16. A method of claim 14, wherein R_1 is fluoro.
 - 17. A method of claim 1, wherein R_2 is nitro.

15

- 18. A method of claim 16, wherein R₃ is H.
- 19. A method of claim 1, wherein R_4 is chloro.
- 20 20. A method of claim 1, wherein R₄ is methyl.
 - 21. A method of claim 1, wherein R_5 is H.
 - 22. A method of claim 1, wherein R₅ is CH₃.

25

- 23. A method of claim 1, wherein X_1 is O or S.
- 24. A method of claim 1, wherein X₁ is NH or NCH₃.
- 30 25. A method of claim 1, wherein X_2 is OH, SH, or NH₂.
 - 26. A method of claim 1, wherein X_2 is NHCH₃ or OH.

10031149.060302 PCT/US00/18346

27. A method of claim 1, wherein said MEK inhibitor has a structure selected from: [5-fluoro-2-(1H-tetrazol-5-yl)-phenyl]-(4-iodo-2-methyl-phenyl)amine; [2,3-difluoro-6-(1H-tetrazol-5-yl)-phenyl]-(4-iodo-2-methyl-phenyl)amine; (4-iodo-2-methyl-phenyl)-[2,3,4-trifluoro-6-(1H-tetrazol-5-yl)-phenyl]-5 amine; [4-bromo-2,3-difluoro-6-(1H-tetrazol-5-yl)-phenyl]-(4-iodo-2-methylphenyl)-amine; [5-fluoro-4-nitro-2-(1H-tetrazol-5-yl)-phenyl]-(4-iodo-2-methylphenyl)-amine; [2-(4,4-dimethyl-4,5-dihydro-oxazol-2-yl)-5-fluoro-phenyl]-(4iodo-2-methyl-phenyl)-amine; [6-(4,4-dimethyl-4,5-dihydro-oxazol-2-yl)-2,3difluoro-phenyl]-(4-iodo-2-methyl-phenyl)-amine; [6-(4,4-dimethyl-4,5-dihydro-10 oxazol-2-yl)-2,3,4-trifluoro-phenyl]-(4-iodo-2-methyl-phenyl)-amine; [4-bromo-6-(4,4-dimethyl-4,5-dihydro-oxazol-2-yl)-2,3-difluoro-phenyl]-(4-iodo-2-methylphenyl)-amine; [2-(4,4-dimethyl-4,5-dihydro-oxazol-2-yl)-5-fluoro-4-nitrophenyl]-(4-iodo-2-methyl-phenyl)-amine; 5-[4-fluoro-2-(4-iodo-2-methyl-15 phenylamino)-phenyl]-[1,3,4]thiadiazol-2-ol; 5-[3,4-difluoro-2-(4-iodo-2-methylphenylamino)-phenyl]-[1,3,4]thiadiazol-2-ol; 5-[3,4,5-trifluoro-2-(4-iodo-2methyl-phenylamino)-phenyl]-[1,3,4]thiadiazol-2-ol; 5-[5-bromo-3,4-difluoro-2-(4-iodo-2-methyl-phenylamino)-phenyl]-[1,3,4]thiadiazol-2-ol; 5-[4-fluoro-2-(4iodo-2-methyl-phenylamino)-5-nitro-phenyl]-[1,3,4]thiadiazol-2-ol; 5-[4-fluoro-20 2-(4-iodo-2-methyl-phenylamino)-phenyl]-[1,3,4]oxadiazol-2-ol; 5-[3,4-difluoro-2-(4-iodo-2-methyl-phenylamino)-phenyl]-[1,3,4]oxadiazol-2-ol; 5-[3,4,5trifluoro-2-(4-iodo-2-methyl-phenylamino)-phenyl]-[1,3,4]oxadiazol-2-ol; 5-[5bromo-3,4-difluoro-2-(4-iodo-2-methyl-phenylamino)-phenyl]-[1,3,4]oxadiazol-2-ol; 5-[4-fluoro-2-(4-iodo-2-methyl-phenylamino)-5-nitro-phenyl]-[1,3,4]oxadiazol-2-ol; 5-[4-fluoro-2-(4-iodo-2-methyl-phenylamino)-phenyl]-4H-25 [1,2,4]triazol-3-ol; 5-[3,4-difluoro-2-(4-iodo-2-methyl-phenylamino)-phenyl]-4H-[1,2,4]triazol-3-ol; 5-[3,4,5-trifluoro-2-(4-iodo-2-methyl-phenylamino)-phenyl]-4H-[1,2,4]triazol-3-ol; 5-[5-bromo-3,4-difluoro-2-(4-iodo-2-methylphenylamino)-phenyl]-4H-[1,2,4]triazol-3-ol; and 5-[4-fluoro-2-(4-iodo-2-30 methyl-phenylamino)-5-nitro-phenyl]-4H-[1,2,4]triazol-3-ol.

PCT/US00/18346

WO 01/05391

28. A method of claim 1, wherein said MEK inhibitor has a structure selected from: 5-[4-fluoro-2-(4-iodo-2-methyl-phenylamino)-phenyl]-[1,3,4]thiadiazol-2-ylamine; 5-[3,4-Difluoro-2-(4-iodo-2-methyl-phenylamino)phenyl]-[1,3,4]thiadiazol-2-ylamine; 5-[3,4,5-Trifluoro-2-(4-iodo-2-methylphenylamino)-phenyl]-[1,3,4]thiadiazol-2-ylamine; 5-[5-bromo-3,4-difluoro-2-5 (4-iodo-2-methyl-phenylamino)-phenyl]-[1,3,4]thiadiazol-2-ylamine; 5-[4-fluoro-2-(4-iodo-2-methyl-phenylamino)-5-nitro-phenyl]-[1,3,4]thiadiazol-2-ylamine; 5-[4-Fluoro-2-(4-iodo-2-methyl-phenylamino)-phenyl]-[1,3,4]oxadiazol-2ylamine; 5-[3,4-difluoro-2-(4-iodo-2-methyl-phenylamino)-phenyl]-10 [1,3,4]oxadiazol-2-ylamine; 5-[3,4,5-trifluoro-2-(4-iodo-2-methyl-phenylamino)phenyl]-[1,3,4]oxadiazol-2-ylamine; 5-[5-bromo-3,4-difluoro-2-(4-iodo-2methyl-phenylamino)-phenyl]-[1,3,4]oxadiazol-2-ylamine; 5-[4-fluoro-2-(4-iodo-2-methyl-phenylamino)-5-nitro-phenyl]-[1,3,4]oxadiazol-2-ylamine; 5-[4-fluoro-2-(4-iodo-2-methyl-phenylamino)-phenyl]-4H-[1,2,4]triazol-3-ylamine; 5-[3,4difluoro-2-(4-iodo-2-methyl-phenylamino)-phenyl]-4H-[1,2,4]triazol-3-ylamine; 15 5-[3,4,5-trifluoro-2-(4-iodo-2-methyl-phenylamino)-phenyl]-4H-[1,2,4]triazol-3ylamine; 5-[5-bromo-3,4-difluoro-2-(4-iodo-2-methyl-phenylamino)-phenyl]-4H-[1,2,4]triazol-3-ylamine; 5-[4-fluoro-2-(4-iodo-2-methyl-phenylamino)-5-nitrophenyl]-4H-[1,2,4]triazol-3-ylamine; 5-[4-fluoro-2-(4-iodo-2-methylphenylamino)-phenyl]-[1,3,4]thiadiazole-2-thiol; 5-[3,4-difluoro-2-(4-iodo-2-20 methyl-phenylamino)-phenyl]-[1,3,4]thiadiazole-2-thiol; 5-[3,4,5-trifluoro-2-(4iodo-2-methyl-phenylamino)-phenyl]-[1,3,4]thiadiazole-2-thiol; 5-[5-bromo-3,4difluoro-2-(4-iodo-2-methyl-phenylamino)-phenyl]-[1,3,4]thiadiazole-2-thiol; 5-[4-fluoro-2-(4-iodo-2-methyl-phenylamino)-5-nitro-phenyl]-[1,3,4]thiadiazole-2-25 thiol: 5-[4-fluoro-2-(4-iodo-2-methyl-phenylamino)-phenyl]-[1,3,4]oxadiazole-2thiol: 5-[3,4-difluoro-2-(4-iodo-2-methyl-phenylamino)-phenyl]-[1,3,4]oxadiazole-2-thiol; 5-[3,4,5-trifluoro-2-(4-iodo-2-methyl-phenylamino)phenyl]-[1,3,4]oxadiazole-2-thiol; 5-[5-bromo-3,4-difluoro-2-(4-iodo-2-methylphenylamino)-phenyl]-[1,3,4]oxadiazole-2-thiol; 5-[4-fluoro-2-(4-iodo-2-methylphenylamino)-5-nitro-phenyl]-[1,3,4]oxadiazole-2-thiol; 5-[4-fluoro-2-(4-iodo-2-30 methyl-phenylamino)-phenyl]-4H-[1,2,4]triazole-3-thiol; 5-[3,4-difluoro-2-(4iodo-2-methyl-phenylamino)-phenyl]-4H-[1,2,4]triazole-3-thiol; 5-[3,4,5-

10031149 060302 PCT/US00/18346

trifluoro-2-(4-iodo-2-methyl-phenylamino)-phenyl]-4H-[1,2,4]triazole-3-thiol; 5-[5-bromo-3,4-difluoro-2-(4-iodo-2-methyl-phenylamino)-phenyl]-4H-[1,2,4]triazole-3-thiol; and 5-[4-fluoro-2-(4-iodo-2-methyl-phenylamino)-5-nitro-phenyl]-4H-[1,2,4]triazole-3-thiol.

5

10

15

20

25

30

A method of claim 1, wherein said MEK inhibitor has a structure 29. selected from: 5-[4-fluoro-2-(4-iodo-2-methyl-phenylamino)-phenyl]-isothiazol-3-ol; 5-[3,4-difluoro-2-(4-iodo-2-methyl-phenylamino)-phenyl]-isothiazol-3-ol; 5-[3,4,5-trifluoro-2-(4-iodo-2-methyl-phenylamino)-phenyl]-isothiazol-3-ol; 5-[5bromo-3,4-difluoro-2-(4-iodo-2-methyl-phenylamino)-phenyl]-isothiazol-3-ol; 5-[4-fluoro-2-(4-iodo-2-methyl-phenylamino)-5-nitro-phenyl]-isothiazol-3-ol; 5-[4fluoro-2-(4-iodo-2-methyl-phenylamino)-phenyl]-isoxazol-3-ol; 5-[3,4-difluoro-2-(4-iodo-2-methyl-phenylamino)-phenyl]-isoxazol-3-ol; 5-[3,4,5-trifluoro-2-(4iodo-2-methyl-phenylamino)-phenyl]-isoxazol-3-ol; 5-[5-bromo-3,4-difluoro-2-(4-iodo-2-methyl-phenylamino)-phenyl]-isoxazol-3-ol; 5-[4-fluoro-2-(4-iodo-2methyl-phenylamino)-5-nitro-phenyl]-isoxazol-3-ol; 5-[4-fluoro-2-(4-iodo-2methyl-phenylamino)-phenyl]-1H-pyrazol-3-ol; 5-[3,4-difluoro-2-(4-iodo-2methyl-phenylamino)-phenyl]-1H-pyrazol-3-ol; 5-[3,4,5-trifluoro-2-(4-iodo-2methyl-phenylamino)-phenyl]-1H-pyrazol-3-ol; 5-[5-bromo-3,4-difluoro-2-(4iodo-2-methyl-phenylamino)-phenyl]-1H-pyrazol-3-ol; 5-[4-fluoro-2-(4-iodo-2methyl-phenylamino)-5-nitro-phenyl]-1H-pyrazol-3-ol; 4-[4-fluoro-2-(4-iodo-2methyl-phenylamino)-phenyl]-isothiazol-3-ol; 4-[3,4-difluoro-2-(4-iodo-2methyl-phenylamino)-phenyl]-isothiazol-3-ol; 4-[3,4,5-trifluoro-2-(4-iodo-2methyl-phenylamino)-phenyl]-isothiazol-3-ol; 4-[5-bromo-3,4-difluoro-2-(4iodo-2-methyl-phenylamino)-phenyl]-isothiazol-3-ol; 4-[4-fluoro-2-(4-iodo-2methyl-phenylamino)-5-nitro-phenyl]-isothiazol-3-ol; 4-[4-fluoro-2-(4-iodo-2methyl-phenylamino)-phenyl]-isoxazol-3-ol; 4-[3,4-difluoro-2-(4-iodo-2-methylphenylamino)-phenyl]-isoxazol-3-ol; 4-[3,4,5-trifluoro-2-(4-iodo-2-methylphenylamino)-phenyl]-isoxazol-3-ol; 4-[5-bromo-3,4-difluoro-2-(4-iodo-2methyl-phenylamino)-phenyl]-isoxazol-3-ol; 4-[4-fluoro-2-(4-iodo-2-methylphenylamino)-5-nitro-phenyl]-isoxazol-3-ol; 4-[4-fluoro-2-(4-iodo-2-methylphenylamino)-phenyl]-1-m thyl-1H-pyrazol-3-ol; 4-[3,4-difluoro-2-(4-iodo-2-

PCT/US00/18346

WO 01/05391

5

methyl-phenylamino)-phenyl]-1-methyl-1H-pyrazol-3-ol; 1-methyl-4-[3,4,5-trifluoro-2-(4-iodo-2-methyl-phenylamino)-phenyl]-1H-pyrazol-3-ol; 4-[5-bromo-3,4-difluoro-2-(4-iodo-2-methyl-phenylamino)-phenyl]-1-methyl-1H-pyrazol-3-ol; and 4-[4-fluoro-2-(4-iodo-2-methyl-phenylamino)-5-nitro-phenyl]-1-methyl-1H-pyrazol-3-ol.

30. A method of claim 1, wherein said MEK inhibitor has a structure selected from: 5-[2-(2-amino-4-iodo-phenylamino)-4-fluoro-phenyl]-1-methyl-1H-[1,2,3]triazol-4-ol; 5-[2-(2-amino-4-iodo-phenylamino)-3,4-difluoro-phenyl]-1-methyl-1H-[1,2,3]triazol-4-ol; 5-[2-(2-amino-4-iodo-phenylamino)-3,4,5-10 trifluoro-phenyl]-1-methyl-1H-[1,2,3]triazol-4-ol; 5-[2-(2-amino-4-iodophenylamino)-5-bromo-3,4-difluoro-phenyl]-1-methyl-1H-[1,2,3]triazol-4-ol; 5-[2-(2-amino-4-iodo-phenylamino)-4-fluoro-5-nitro-phenyl]-1-methyl-1H-[1,2,3]triazol-4-ol; 5-[4-fluoro-2-(4-iodo-2-methyl-phenylamino)-phenyl]-3-15 methyl-3H-[1,2,3]triazol-4-ol; 5-[3,4-difluoro-2-(4-iodo-2-methyl-phenylamino)phenyl]-3-methyl-3H-[1,2,3]triazol-4-ol; 3-methyl-5-[3,4,5-trifluoro-2-(4-iodo-2methyl-phenylamino)-phenyl]-3H-[1,2,3]triazol-4-ol; 5-[5-bromo-3,4-difluoro-2-(4-iodo-2-methyl-phenylamino)-phenyl]-3-methyl-3H-[1,2,3]triazol-4-ol; 5-[4fluoro-2-(4-iodo-2-methyl-phenylamino)-5-nitro-phenyl]-3-methyl-3H-[1,2,3]triazol-4-ol; 4-[4-fluoro-2-(4-iodo-2-methyl-phenylamino)-phenyl]-2-20 methyl-2H-pyrazol-3-ol; 4-[3,4-difluoro-2-(4-iodo-2-methyl-phenylamino)phenyl]-2-methyl-2H-pyrazol-3-ol; 2-methyl-4-[3,4,5-trifluoro-2-(4-iodo-2methyl-phenylamino)-phenyll-2H-pyrazol-3-ol; 4-[5-bromo-3,4-difluoro-2-(4iodo-2-methyl-phenylamino)-phenyl]-2-methyl-2H-pyrazol-3-ol; 4-[4-fluoro-2-25 (4-iodo-2-methyl-phenylamino)-5-nitro-phenyl]-2-methyl-2H-pyrazol-3-ol; 1-[4fluoro-2-(4-iodo-2-methyl-phenylamino)-phenyll-4-methyl-1,4-dihydro-tetrazol-5-one; 1-[3,4-difluoro-2-(4-iodo-2-methyl-phenylamino)-phenyl]-4-methyl-1,4dihydro-tetrazol-5-one; 1-methyl-4-[3,4,5-trifluoro-2-(4-iodo-2-methylphenylamino)-phenyl]-1,4-dihydro-tetrazol-5-one; 1-[5-bromo-3,4-difluoro-2-(4-iodo-2-methyl-phenylamino)-phenyl]-4-methyl-1,4-dihydro-tetrazol-5-one; 1-30 [4-fluoro-2-(4-iodo-2-methyl-phenylamino)-5-nitro-phenyl]-4-methyl-1,4dihydro-tetrazol-5-one; 1-[4-fluoro-2-(4-iodo-2-methyl-phenylamino)-phenyl]-

5

30

10031149 "06030E PCT/US00/18346

1H-[1,2,3]triazol-4-ol; 1-[3,4-difluoro-2-(4-iodo-2-methyl-phenylamino)-phenyl]-1H-[1,2,3]triazol-4-ol; 1-[3,4,5-trifluoro-2-(4-iodo-2-methyl-phenylamino)-phenyl]-1H-[1,2,3]triazol-4-ol; 1-[5-bromo-3,4-difluoro-2-(4-iodo-2-methyl-phenylamino)-phenyl]-1H-[1,2,3]triazol-4-ol; and 1-[4-fluoro-2-(4-iodo-2-methyl-phenylamino)-5-nitro-phenyl]-1H-[1,2,3]triazol-4-ol.

- 31. The method of claim 1, wherein said MEK inhibitor has a structure selected from: 3-[4-fluoro-2-(4-iodo-2-methyl-phenylamino)-phenyl]-2Hisoxazol-5-one; 3[3,4-difluoro-2-(4-iodo-2-methyl-phenylamino)-phenyl]-2Hisoxazol-5-one; 3-[3,4,5-trifluoro-2-(4-iodo-2-methyl-phenylamino)-phenyl]-2H-10 isoxazol-5-one; 3-[5-bromo-3,4-difluoro-2-(4-iodo-2-methyl-phenylamino)phenyl]-2H-isoxazol-5-one; 3-[4-fluoro-2-(4-iodo-2-methyl-phenylamino)-5nitro-phenyl]-2H-isoxazol-5-one; [5-fluoro-2-(2-oxo-2,3-dihydro-2l>4 -[1,2,3,5]oxathiadiazol-4-yl)-phenyl]-(4-iodo-2-methyl-phenyl)-amine; [2,3difluoro-6-(2-oxo-2,3-dihydro-2I>4 -[1,2,3,5]oxathiadiazol-4-yI)-phenyI]-(4-15 iodo-2-methyl-phenyl)-amine; (4-lodo-2-methyl-phenyl)-[2,3,4-trifluoro-6-(2oxo-2,3-dihydro-2l>4 -[1,2,3,5]oxathiadiazol-4-yl)-phenyl]-amine; [4-bromo-2,3-difluoro-6-(2-oxo-2,3-dihydro-2l>4_-[1,2,3,5]oxathiadiazol-4-yl)-phenyl]-(4iodo-2-methyl-phenyl)-amine; [5-fluoro-4-nitro-2-(2-oxo-2,3-dihydro-2l>4_-[1,2,3,5]oxathiadiazol-4-yl)-phenyl]-(4-iodo-2-methyl-phenyl)-amine; 4-[4-20 fluoro-2-(4-iodo-2-methyl-phenylamino)-phenyl]-4H-isoxazol-5-one; 4-[3,4difluoro-2-(4-iodo-2-methyl-phenylamino)-phenyl]-4H-isoxazol-5-one; 4-[3,4,5trifluoro-2-(4-iodo-2-methyl-phenylamino)-phenyl]-4H-isoxazol-5-one; 4-[5bromo-3,4-difluoro-2-(4-iodo-2-methyl-phenylamino)-phenyl]-4H-isoxazol-5one; and 4-[4-fluoro-2-(4-iodo-2-methyl-phenylamino)-5-nitro-phenyl]-4H-25 isoxazol-5-one.
 - 32. The method of claim 1, wherein said MEK inhibitor has a structure selected from: 2,4-bis-(2-chloro-4-iodo-phenylamino)-3fluoro5-nitro-benzoic acid; 5-[3,4difluoro-2-(4-iodo-2-methyl-phenylamino)-phenyl]-[1,3,4]oxadiazol-2-ylamine; 5-[4-fluoro-2-(4-iodo-2-methyl-phenylamino)-phenyl]-[1,3,4]oxadiazol-2-ol; (2,3-difluoro-6-[1,3,4]oxadiazol-2-yl-phenyl)(-(4-iodo-2-methyl-phenyl))-[1,3,4]oxadiazol-2-yl-phenyl)(-(4-iodo-2-methyl-phenyl))-[1,3,4]oxadiazol-2-yl-phenyl)(-(4-iodo-2-methyl-phenyl))-[1,3,4]oxadiazol-2-yl-phenyl)(-(4-iodo-2-methyl-phenyl))-[1,3,4]oxadiazol-2-yl-phenyl)(-(4-iodo-2-methyl-phenyl))-[1,3,4]oxadiazol-2-yl-phenyl)(-(4-iodo-2-methyl-phenyl))-[1,3,4]oxadiazol-2-yl-phenyl)-[1,3,4]oxadiazol-2-yl-phenyl)-[1,3,4]oxadiazol-2-yl-phenyl)-[1,3,4]oxadiazol-2-yl-phenyl)-[1,3,4]oxadiazol-2-yl-phenyl)-[1,3,4]oxadiazol-2-yl-phenyl)-[1,3,4]oxadiazol-2-yl-phenyl)-[1,3,4]oxadiazol-2-yl-phenyl)-[1,3,4]oxadiazol-2-yl-phenyl)-[1,3,4]oxadiazol-2-yl-phenyl)-[1,3,4]oxadiazol-2-yl-phenyl-phenyl-phenyl-phenyl-phenyl-phenyl-phenyl-phenyl-phenyl-phenyl-phenyl-phenyl-phenyl-phenyl-phenyl-phenyl-phenyl-phenyl-phenyl-phenyl-phenyl-phenyl-phenyl-phenyl-phenyl-phenyl-phenyl-phenyl-phenyl-phenyl-phenyl-phenyl-phenyl-phenyl-phenyl-phenyl-phenyl-phenyl-phenyl-phenyl-phenyl-phenyl-phenyl-phenyl-phenyl-phenyl-phenyl-phenyl-phenyl-phenyl-phenyl-phenyl-phenyl-phenyl-phenyl-phenyl-phenyl-phenyl-phenyl-phenyl-phenyl-phenyl-phenyl-phenyl-phenyl-phenyl-phenyl-phenyl-phenyl-phenyl-phenyl-phenyl-phenyl-phenyl-phenyl-phenyl-phenyl-phenyl-phenyl-phenyl-phenyl-phenyl-phenyl-phenyl-phenyl-phenyl-phenyl-phenyl-phenyl-phenyl-phenyl-phenyl-phenyl-phenyl-phenyl-phenyl-phenyl-phenyl-phenyl-phenyl-phenyl-phenyl-phenyl-phenyl-phenyl-phenyl-phenyl-phenyl-phenyl-phenyl-phenyl-phenyl-phenyl-phenyl-phenyl-phenyl-phenyl-phenyl-phenyl-phenyl-phenyl-phenyl-phenyl-phenyl-phenyl-phenyl-phenyl-phenyl-phenyl-phenyl-phenyl-phenyl-phenyl-phenyl-phenyl-phenyl-phenyl-phenyl-phenyl-phenyl-phenyl-phenyl-phenyl-phenyl-phenyl-phenyl-phenyl-phenyl-phenyl-phenyl-phenyl

WO 01/05391 PCT/US00/18346

methyl-phenyl)-amine; 5-[3,4-difluoro-2-(4-iodo-2-methyl-phenylamino)-phenyl]-[1,3,4]oxadiazole-2-thiol; 5-[3,4difluoro-2-(4-iodo-2-methyl-phenyl]-4*H*-[1,2,4]triazole-3-ylamine; and 5-[3,4-difluoro-2-(4-iodo-2-methyl-phenylamino)-phenyl]-4*H*-[1,2,4]triazole-3-thiol.

5

33. The method of claim 1, wherein said MEK inhibitor has the structure: 2,4-bis-(2-chloro-4-iodo-phenylamino)-3-fluoro-5-nitro-benzoic acid.